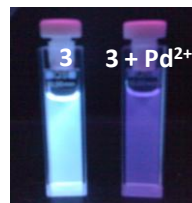
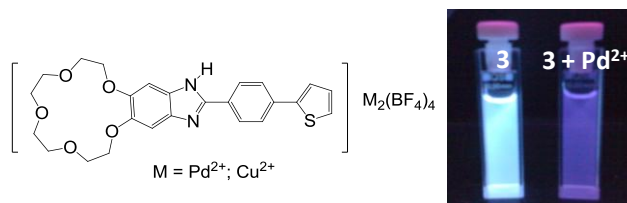


GRAPHICAL ABSTRACT

Imidazo-benzo-15-crown-5 ethers bearing arylthienyl and bithienyl moieties as novel fluorescent chemosensors for Pd^{2+} and Cu^{2+}

Rosa M. F. Batista, Elisabete Oliveira, Susana P. G. Costa,¹ Carlos Lodeiro and M. Manuela M. Raposo



Imidazo-benzo-15-crown-5 ethers bearing arylthienyl and bithienyl moieties as novel fluorescent chemosensors for Pd²⁺ and Cu²⁺

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Abstract- Novel fluorescent ionophores bearing imidazo-arylthienyl or imidazo-bithienyl π -conjugated bridges functionalized with one or two fused benzo-15-crown-5 ethers as receptor units are reported. The sensing ability of the compounds in the presence of metallic cations (Li⁺, Na⁺, K⁺, Ca²⁺, Zn²⁺, Cu²⁺, Ni²⁺, Pd²⁺ and Hg²⁺) and fluoride ion was studied in MeCN/DMSO solutions by absorption and emission spectroscopy. The experimental results indicate that all compounds could act as selective fluorimetric sensors for Cu²⁺ and Pd²⁺ and also for the fluoride ion, in the case of the *bis*-substituted crown ether derivatives.

Keywords: Crown ethers, Arylthiophene, Bithiophene, Imidazole, Fluorescent chemosensors, CHEQ effect, Pd²⁺, Cu²⁺, F⁻.

1. Introduction

The development of artificial receptors for the sensing and recognition of environmentally and biologically important ionic species is currently of great interest. Highly selective anion¹ or cation² sensing is imperative for many areas, including environmental, biological, clinical, and waste management applications.

The potential for progress in the development of fluorescent sensors can be explained by the distinct advantages offered by fluorescence detection in terms of sensitivity, selectivity, response time, local observation, etc. Typically, an auxochrome generating the fluorescence signal is combined with an analyte-responsive receptor *via* a saturated or unsaturated spacer. The sensing properties of such systems depend on the receptor-controlled (chemical) selectivity and the analyte-mediated (spectroscopic) signaling selectivity as well as sensitivity. Suitable fluorescent reporters must efficiently transduce a binding event into a measurable fluorescence signal, thereby taking advantage of the intrinsic selectivity of communication of fluorescence *via* two experimental parameters, excitation and emission wavelength, and its high sensitivity.^{1c-d,3}

Crown ether derivatives occupy a special position among receptors and are widely used in the design of new chemosensors based on their unique ability to coordinate the cations of alkaline metals, their fairly high selectivity and accessibility. In addition to alkaline metals, crown ethers are also effective complexing reagents for the cations of alkaline-earth metal ions and transition metal ions.⁴ On the other hand, hydrogen-bond donors such as pyrrole/calixpyrrole, (thio)urea, dipyrrolylquinoxalines, indolocarbazoles, and (benzo)imidazole usually act as anion binding sites and the acidity of the NH proton of the imidazole can be tuned by changing the electronic properties of the imidazole substituents.⁵ Recently, 2,4,5-triaryl(heteroaryl)-imidazole-based chromophores have received increasing attention due to their interesting optical properties. For example, one of the most interesting applications are as fluorescent chemosensors.^{6a-c} The development of chemosensors for the sensing of transition metal ions is one of the most active research fields with great potential for environmental and physiological applications, in particular Cu²⁺-sensitive systems.⁷ Even though great achievement in the field of colorimetric and/or fluorescent chemosensors for Cu²⁺ has been obtained,⁸ there is still a demand to develop new indicators with improved properties, especially fluorescent sensors with high efficiency in the visible spectra and with specific selectivity towards Cu²⁺ over other competitive metal ions. Also, due to the widespread application of Pd²⁺ as catalyst in organic synthesis is important the development of novel systems for its easy detection specially because a high level of residual palladium (typically 300-2000 ppm) is often found in the resultant products, which thus may be a health hazard.⁹

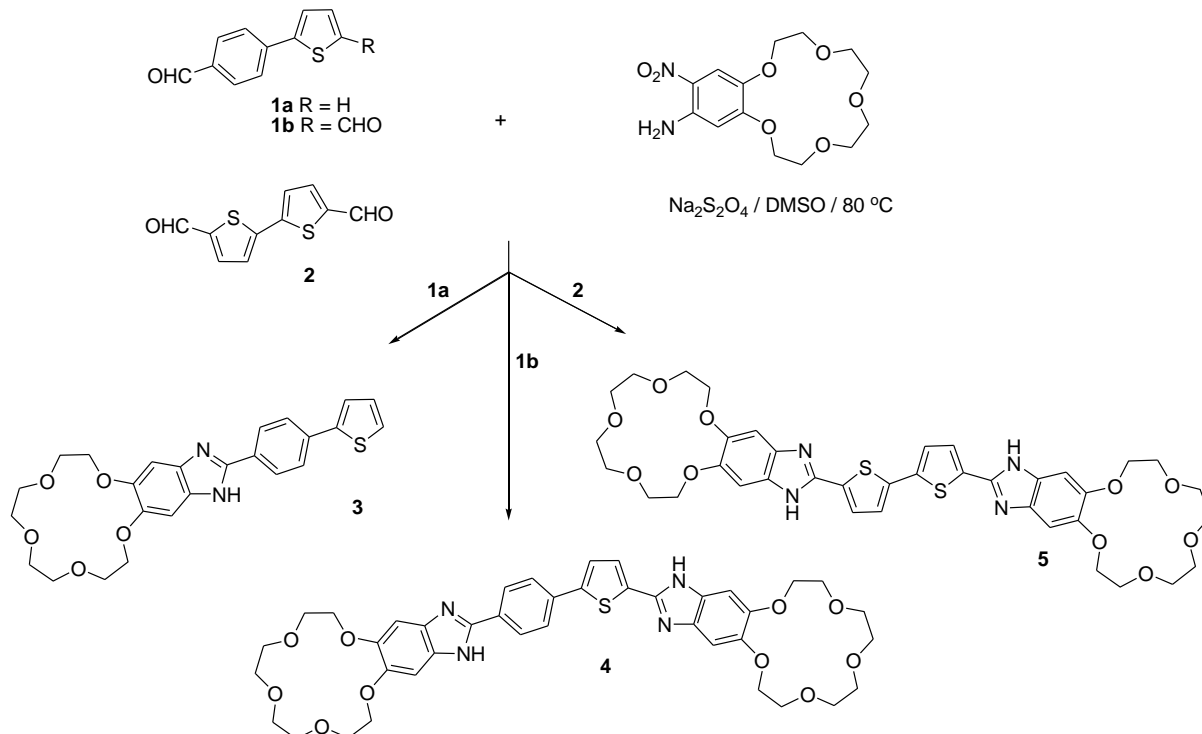
We have demonstrated that oligothiophene and arylthiophene derivatives, electronically connected to recognition sites, are efficient π -conjugated bridges for the fluorimetric and/colorimetric sensing of certain anions (*e.g.*, F⁻, CN⁻) and cations (H⁺, Na⁺, Pd²⁺, Cu²⁺,

Zn^{2+} , Hg^{2+} , Ni^{2+}).^{5d,6c,10} Following our current interests on colorimetric and fluorimetric chemosensors provided with heterocyclic moieties,^{5d,6c,10,11} three novel heterocyclic systems containing arylthiophene (**3-4**) or bithiophene (**5**) bridges and one or two imidazo-benzocrown ether binding moieties were designed and synthesized.

2. Results and discussion

2.1. Synthesis

Recently the application of a mild and versatile method for 2-benzimidazole synthesis, was reported through a one step reaction involving $\text{Na}_2\text{S}_2\text{O}_4$ reduction of *o*-nitroanilines in the presence of pyridyl and quinolyl aldehydes¹² in DMSO. To the best of our knowledge, this is the first time that this synthetic methodology is applied to arylthiophene and bithiophene aldehydes and an *o*-nitroaminobenzocrown ether derivative. Therefore, formyl-arylthiophenes **1a-b**, diformyl-bithiophene **2** and 4'-amino-5'-nitrobenzo-15-crown-5 ether were used in the synthesis of imidazo-benzocrown ethers **3-5**, using the synthetic methodology described above in order to evaluate the influence of the different π -conjugated bridges on the optical and sensing properties of the new compounds (Scheme 1).



Scheme 1. Synthesis of imidazo-benzocrown ether derivatives **3-5**.

Compounds **3-5** were obtained in moderate to good yields (53-78%) and their structures were unambiguously confirmed by their analytical and spectral data (Table 1). Aldehyde precursors were commercially available (**1a**) or synthesised as reported previously by some of us (**1b**, **2**).^{6d,13}

Table 1. Yields, IR, ¹H NMR, UV-vis absorption and fluorescence data for imidazo-benzocrown ethers **3-5**.

Cpd.	Yield (%)	IR ν (cm ⁻¹)	δ_{H} (ppm) ^c	UV/Vis		Fluorescence	
				λ_{max} (nm) ^d	λ_{em} (nm) ^d	Stokes' shift (nm)	Φ_{F}
3	64	3443 ^a	----	360	447	87	0.76
4	78	3449 ^a	13.45	395	500	105	0.50
5	53	3451 ^b	12.76	415	568	135	0.10

^a For the NH stretching band (recorded in nujol).

^b For the NH stretching band (recorded in KBr).

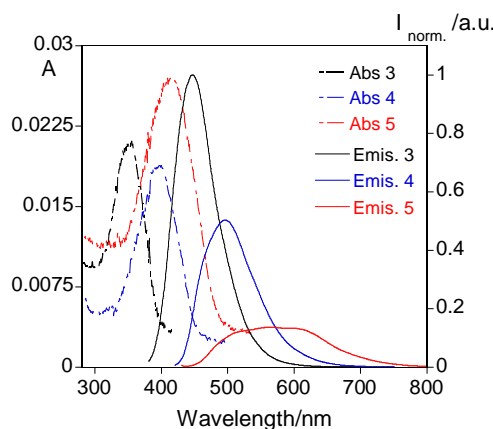
^c For the NH proton of the imidazole ring (in DMSO-d₆).

^d in MeCN/DMSO (99:1) solution.

2.2. Photophysical study of imidazo-benzocrown ether derivatives **3-5**.

The electronic absorption and emission spectra of compounds **3-5** were obtained in MeCN/DMSO (99:1) solution (Table 1). The comparison of the absorption maxima of compounds **3** and **4** reveals the influence of the introduction of a second imidazo-benzocrown ether moiety as the longest wavelength transition was shifted from 360 to 395 nm, due to more extensive electron delocalization. When comparing compounds **4** and **5**, which differ in the electronic nature of the π -conjugated bridge, it can be seen that compound **5** shows a higher intramolecular charge transfer efficiency due to the substitution of a phenyl for a thiophene ring. The same trend was observed in the emission spectra for related compounds.¹⁴ The study of the fluorescence properties of compounds **3-5** showed that they are strongly emissive, with relative fluorescence quantum yields ranging from 0.10 to 0.76 (Figure 1).

Figure 1. Absorption and emission spectra of compounds **3-5** in MeCN/DMSO (99:1), T= 298 K, [**3**] = 5.0×10^{-7} M, [**4**] = 5.5×10^{-7} M, [**5**] = 4.3×10^{-7} M. ($\lambda_{\text{exc3}} = 360$ nm, $\lambda_{\text{exc4}} = 395$ nm; $\lambda_{\text{exc5}} = 415$ nm).



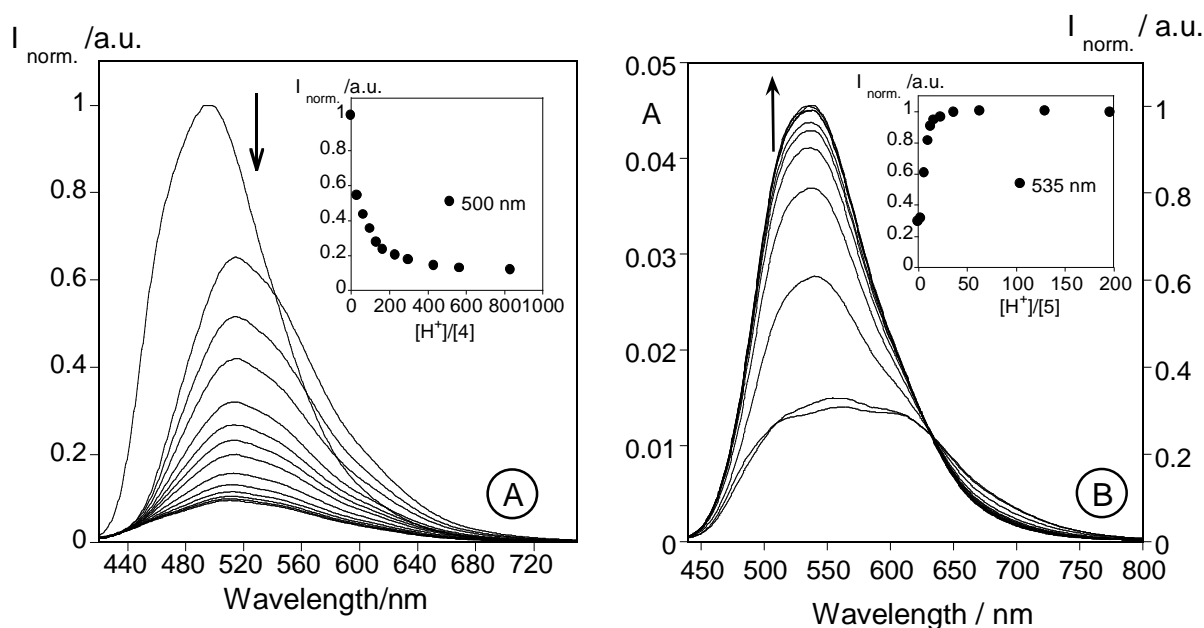
Our results revealed a decrease in the relative fluorescence quantum yield when a second imidazo-benzocrown was introduced (**3** compared to **4**), due to the azomethine nitrogens that contribute to the heavy atom effect concomitant with the increased degree of conjugation,¹⁵ or when a thiophene replaced a phenyl ring (**4** compared to **5**), with the heavy atom induced spin-orbit coupling by the sulphur atoms giving rise to a very efficient intersystem crossing mechanism, that lowers the emission.^{14,16} In the latter case, the decrease was larger pointing to the better quenching process produced by the sulphur atom present in the thiophene when compared to the nitrogen present in the imidazole.

2.3. Spectrophotometric and spectrofluorimetric titrations with proton, metallic cations and fluoride anion

Compounds **3-5** were also evaluated as chemosensors in the presence of H^+ and several metal cations (Li^+ , Na^+ , K^+ , Ca^{2+} , Zn^{2+} , Cu^{2+} , Ni^{2+} , Pd^{2+} , and Hg^{2+}) in MeCN/DMSO (99:1) solutions. Having in mind our recent results concerning the sensing properties of imidazo-anthraquinone derivatives bearing thiophene moieties,^{5d} compounds **3-5** were also studied as chemosensors for the F^- anion. It is known the acidity of the NH proton of the imidazole ring can also be tuned by changing the electronic properties of the π -conjugated bridge and our results for the signal of NH appearing downfield in the ^1H NMR spectra indicated high acidity and strong hydrogen bonding ability.

Due to the presence of protonable imidazo units, all reported compounds were studied in the presence of H^+ , by titration with a solution of methanesulphonic acid in MeCN/DMSO (99:1). In general, the absorption spectra of all compounds were slightly affected, showing a red-shift of 10 nm. Regarding the emission spectra, for **3** the intensity of the emission was practically unaffected upon addition of 50 equivalents of acid, whereas addition of the same amount of acid to **4** induced a 50% quenching, and **5** showed a remarkable 70% increase in fluorescence intensity and a 20 nm blue shift. This effect can be probably due to the different aromatic bridge present in **4** and **5**, bearing spacers of different planarity between both imidazo-benzo units.¹⁸ At the same time, the protonation at the nitrogen of the imidazo group resulted in a partial quenching.^{18b} (Figure 2).

Figure 2. Spectrofluorimetric titration of compounds **4** (A) and **5** (B) in MeCN/DMSO (99:1) solutions as a function of added CH_4O_3S . The insets show the normalized fluorescence intensity at 500 nm and 535 nm, respectively ($[4] = 4.30 \times 10^{-7}$ M, $\lambda_{exc} = 395$ nm; $[5] = 5.10 \times 10^{-7}$ M, $\lambda_{exc} = 415$ nm).

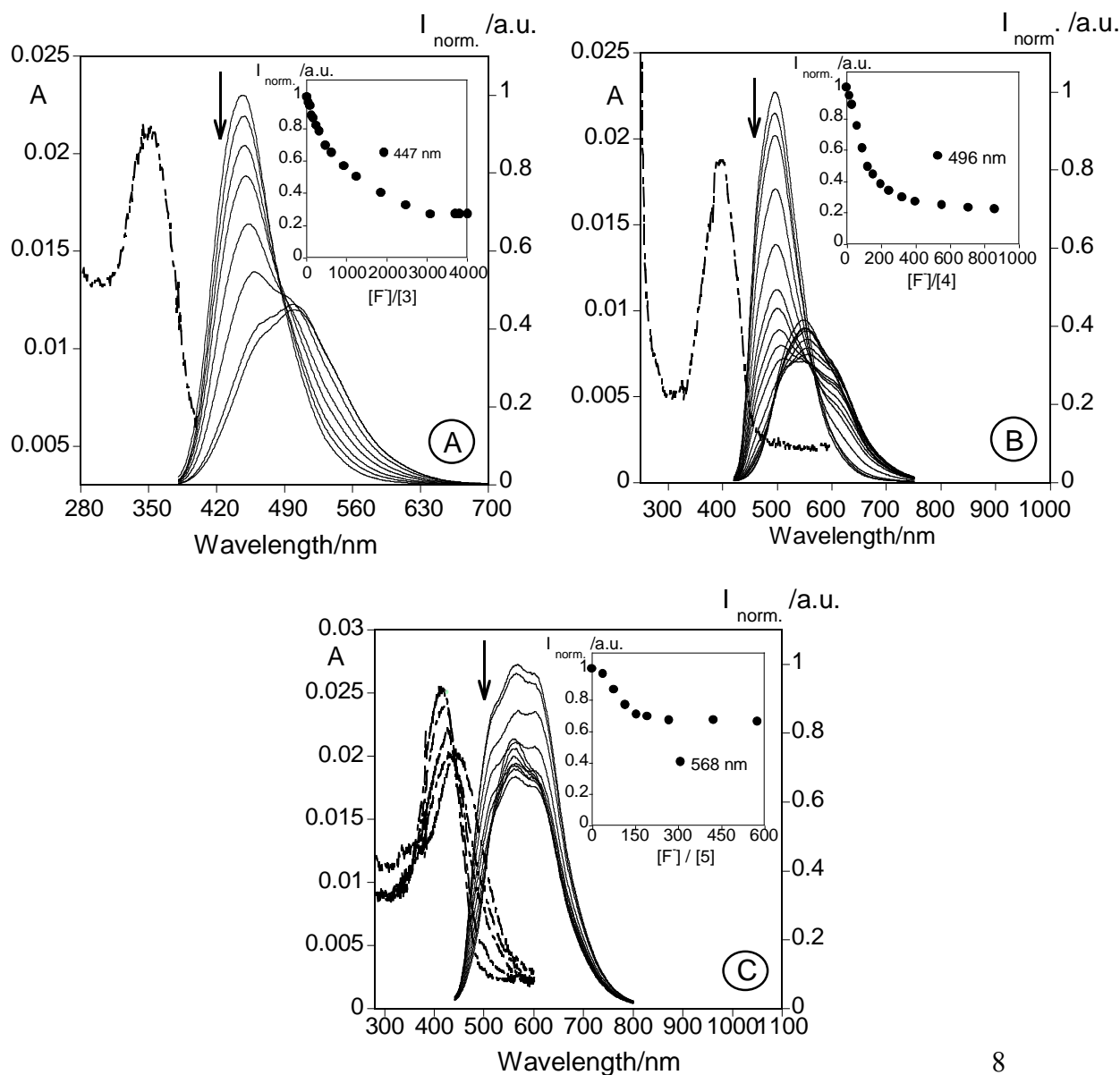


For all compounds, the fluorescence intensity was quenched upon addition of fluoride ion. For compound **3**, 4000 equivalents were necessary to reach a plateau, while compounds **4** and **5** required the addition of 200 and 260 equivalents, respectively. Compound **4** appears to be the most acidic, a fact that is also confirmed by the 1H NMR NH signal appearing downfield (when compared to **5**). A blue shift in the emission band was observed at the same time for all

the compounds. (Figure 3). This quenching effect can be understood as a PET effect from the lone pair of electrons located at the deprotonated imidazole ring.

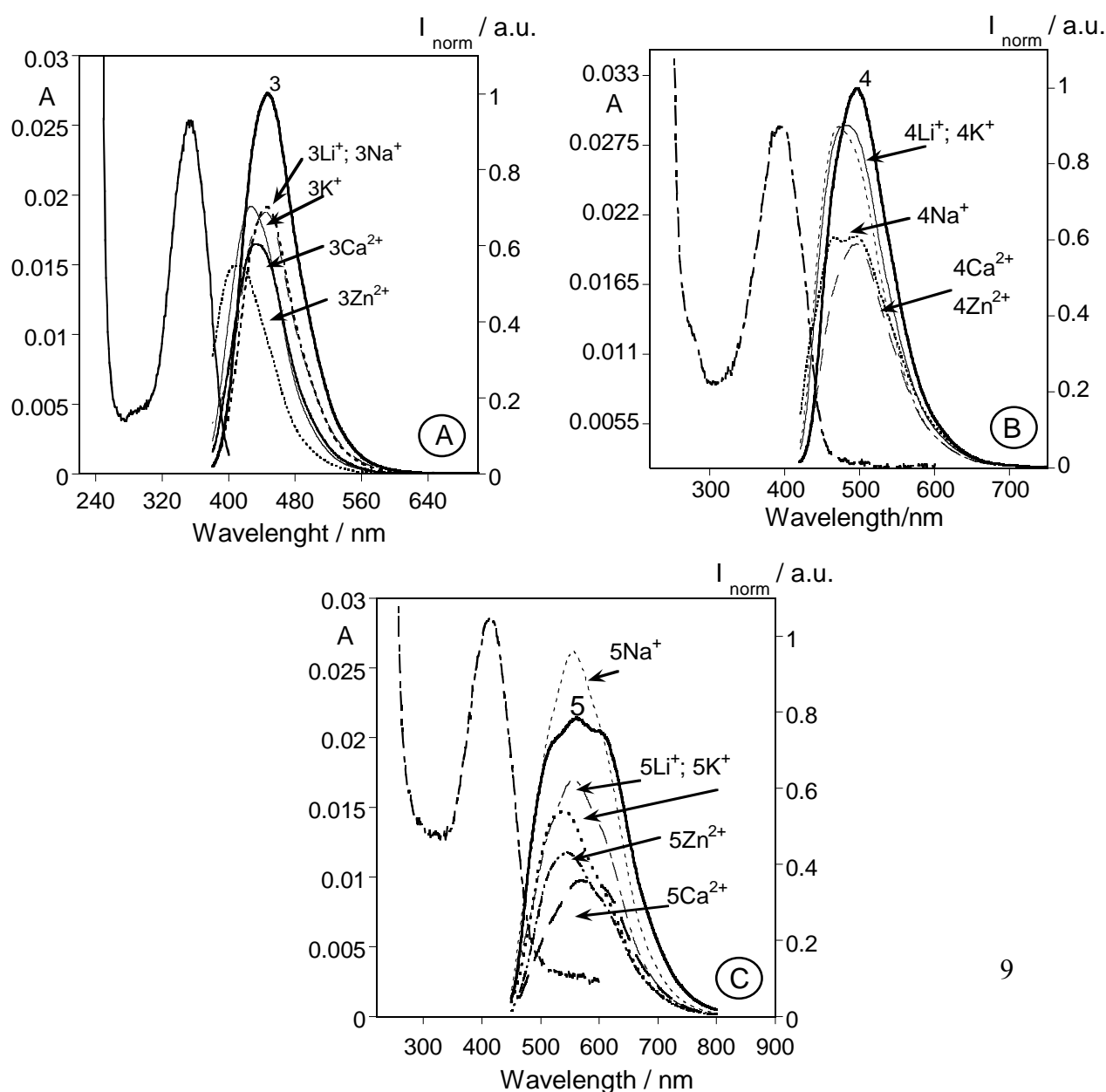
The sensing ability of compounds **3** to **5** was evaluated through interaction with metal ions such as alkaline (Na^+ , Li^+ and K^+), alkaline-earth (Ca^{2+}), and transition and post-transition metal cations (Zn^{2+} , Cu^{2+} , Ni^{2+} , Pd^{2+} and Hg^{2+}) in MeCN/DMSO (99:1) by spectrophotometric and spectrofluorimetric titrations.

Figure 3. Absorption and fluorescence spectra of MeCN/DMSO (99:1) solutions of **3** (A), **4** (B) and **5** (C) and fluorescence titration as a function of Bu_4NF added. The insets show the normalized fluorescence intensity at 447 nm (**3**), 496 nm (**4**) and 568 nm (**5**) ($[\mathbf{3}] = 5.13 \times 10^{-7}$ M, $\lambda_{\text{exc}} = 360$ nm; $[\mathbf{4}] = 4.30 \times 10^{-7}$ M, $\lambda_{\text{exc}} = 395$ nm). $[\mathbf{5}] = 5.10 \times 10^{-7}$ M, $\lambda_{\text{exc}} = 415$ nm).



The addition of up to 2000 equivalents of Na^+ , Li^+ and K^+ metal ions practically does not induce spectral changes, producing only a 10% quenching of the fluorescence emission. However, in compound **5** the addition of the same amount of Na^+ resulted in a blue-shift in the emission band and an intensity increase of 20% reaching a plateau (Figure 4). It is well known that the benzo-15-crown-5-ether quelating unit is used specifically for Na^+ sensing. However, our results suggest that the high delocalized charge presented in these conjugated systems could modulate the interaction with the crown ether resulting in small changes in the ground state, as well as in the excited state.

Figure 4. Absorption and fluorescence spectra of MeCN/DMSO (99:1) solutions of **3** (A), **4** (B) and **5** (C) after addition of 2000 equivalents of NaBF_4 ; LiBF_4 ; KBF_4 ; $\text{Ca}(\text{CF}_3\text{SO}_3)_2$ and $\text{Zn}(\text{CF}_3\text{SO}_3)_2$ in CH_3CN . ($[\mathbf{3}] = 5.13 \times 10^{-7} \text{ M}$, $\lambda_{\text{exc}} = 360 \text{ nm}$; $[\mathbf{4}] = 4.30 \times 10^{-7} \text{ M}$, $\lambda_{\text{exc}} = 395 \text{ nm}$; $[\mathbf{5}] = 5.10 \times 10^{-7} \text{ M}$, $\lambda_{\text{exc}} = 415 \text{ nm}$).

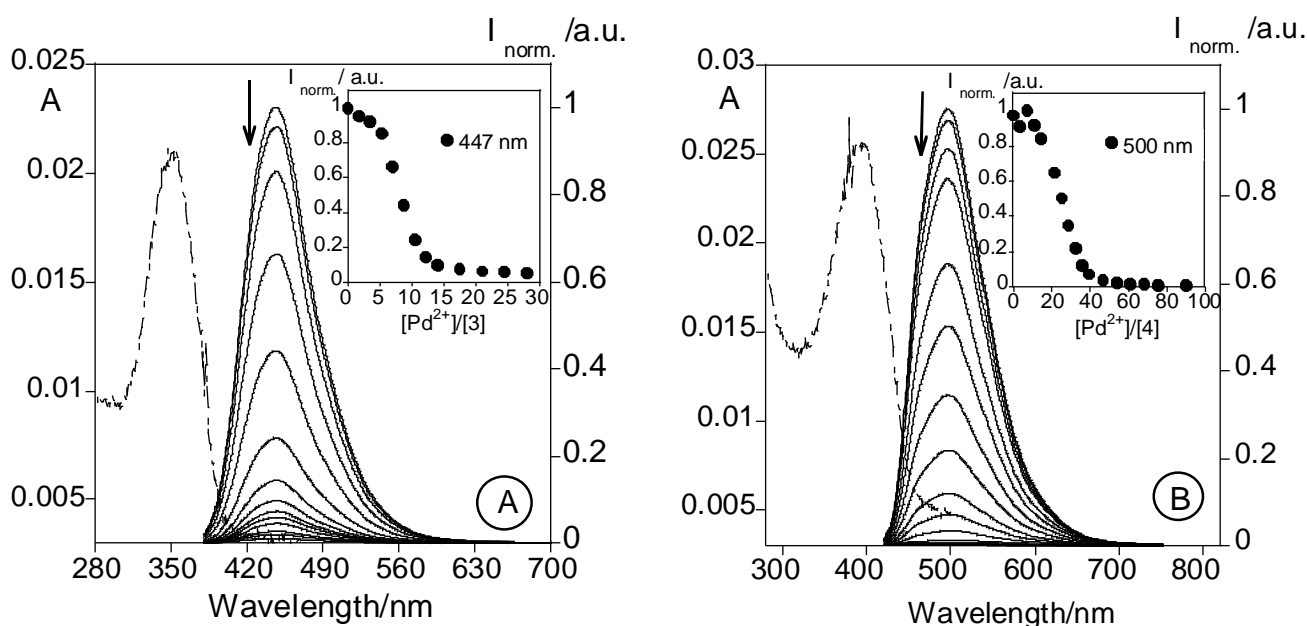


In general, the interaction of compounds **3-5** with Zn^{2+} , Ni^{2+} and Hg^{2+} show a very small blue shift, between 10 and 15 nm, in the absorption band. In the case of Zn^{2+} that normally produces a CHEF (chelation enhancement of fluorescence) effect with polyamine systems, it was observed a 40 to 50% quenching of emission in all cases. Similar findings had been published for compounds bearing a coordinative aromatic nitrogen atom.¹⁷ Concerning Ni^{2+} and Hg^{2+} , it was observed the same quenching effect of about 50% with compound **4**.

The strongest interaction observed for the new compounds reported was found in the presence of Pd^{2+} and Cu^{2+} .

A CHEQ (chelation enhancement of the quenching) effect in the fluorescence emission for these both metals was observed (Figure 5 and Figure 6). Moreover, a blue shift was observed when Cu^{2+} was added to ligand **5** (Figure 6). The stronger interaction was detected for compound **4** in the presence of Cu^{2+} , where only 25 equivalents are enough to quench the emission. In the case of compounds **3** and **5**, up to 40 and 60 equivalents were necessary, respectively.

Figure 5. Absorption spectra of MeCN-DMSO (99:1) solutions of **3** (A), **4** (B) and **5** (C) and fluorescence titration as a function of added $\text{Pd}(\text{CH}_3\text{CN})_4\text{BF}_4$. The insets show the normalized fluorescence intensity at 447 nm (**3**), 500 nm (**4**) and 568 nm (**5**). ($T = 298 \text{ K}$; $[\mathbf{3}] = 5.13 \cdot 10^{-7} \text{ M}$, $\lambda_{\text{exc}} = 360 \text{ nm}$; $[\mathbf{4}] = 5.10 \cdot 10^{-7} \text{ M}$, $\lambda_{\text{exc}} = 395 \text{ nm}$; $[\mathbf{5}] = 4.30 \cdot 10^{-7} \text{ M}$, $\lambda_{\text{exc}} = 415 \text{ nm}$).



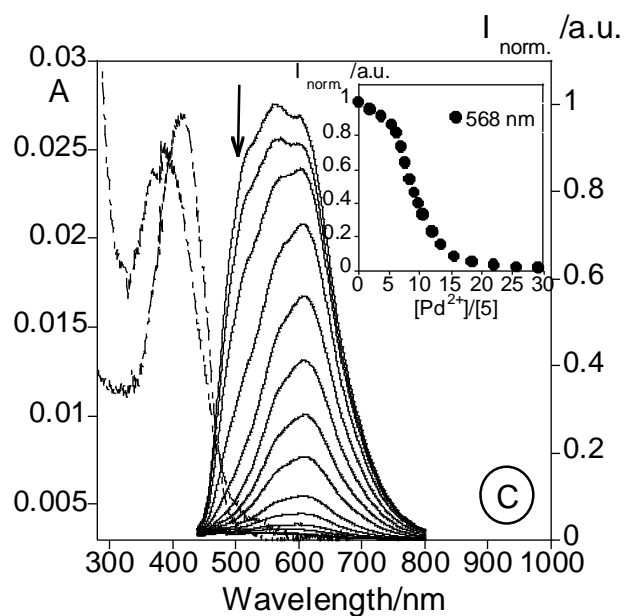
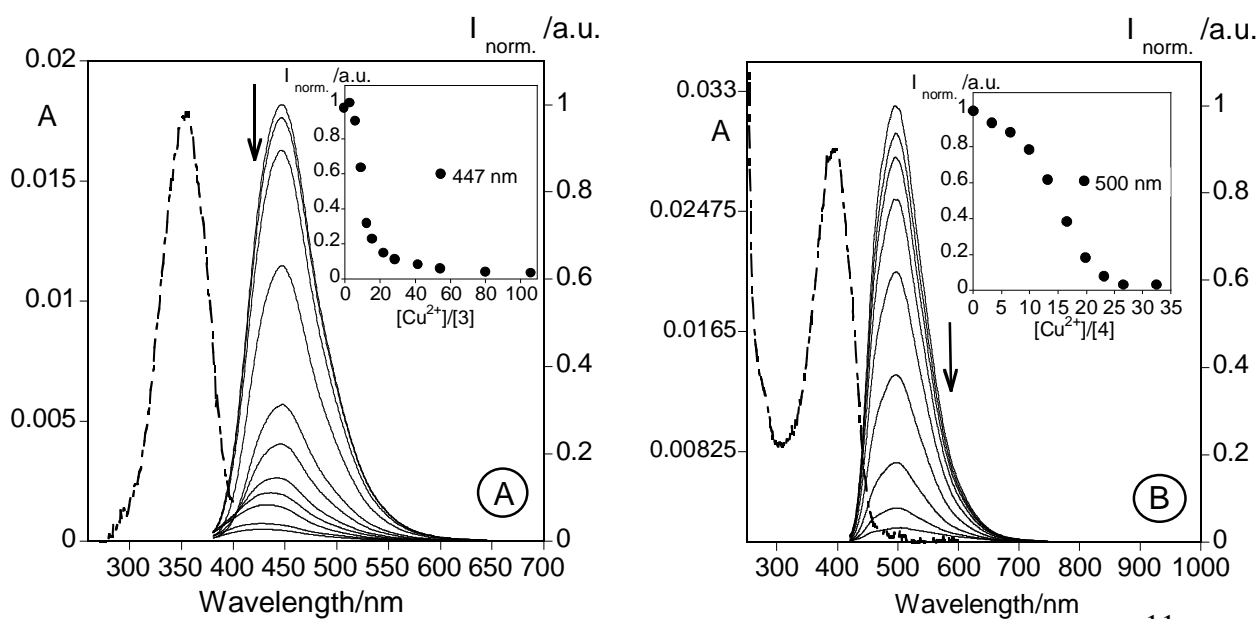
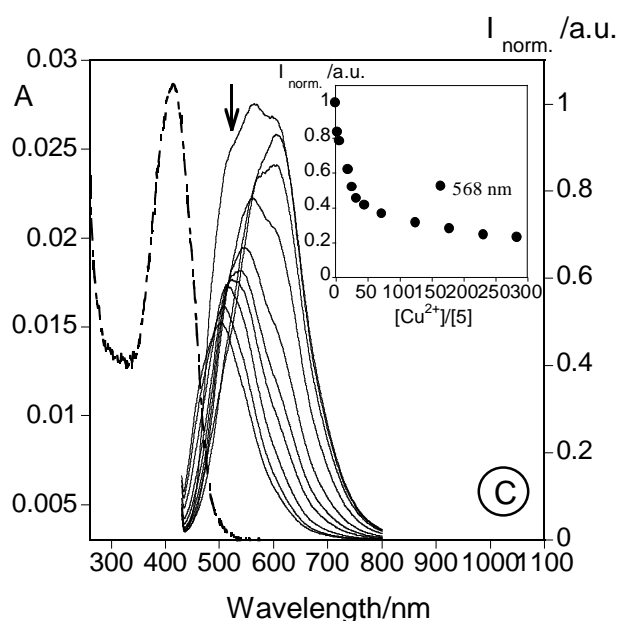


Figure 6. Absorption and fluorescence spectra of MeCN/DMSO (99:1) solutions of **3** (A), **4** (B) and **5** (C) as a function of added $\text{Cu}(\text{CF}_3\text{SO}_3)_2$. The insets show the normalized fluorescence intensity at 447 nm (**3**), 500 nm (**4**) and 568 nm (**5**). ($[\text{3}] = 5.13 \times 10^{-7} \text{ M}$, $\lambda_{\text{exc}} = 360 \text{ nm}$; $[\text{4}] = 4.30 \times 10^{-7} \text{ M}$, $\lambda_{\text{exc}} = 395 \text{ nm}$, $[\text{5}] = 5.10 \times 10^{-7} \text{ M}$, $\lambda_{\text{exc}} = 415 \text{ nm}$).



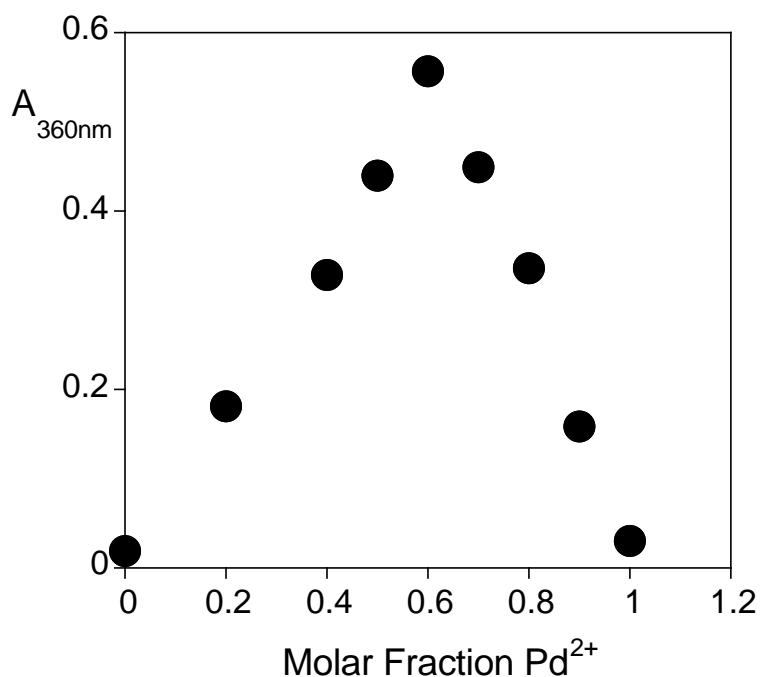


Insets of Figure 5 show the metal ion titration followed at the maximum of the emission band. Inspection of these figures shows that compound **3** and **5** are totally quenched upon the addition of 15 equivalents of Pd^{2+} , while in compound **4**, 40 equivalents to obtain the same quenching effect were necessary. In summary, compounds **3** and **5** appear to be quenched more efficiently by Pd^{2+} while compound **4** by Cu^{2+} . This result turns out from the involvement of the nitrogen atom from the imidazole ring and the sulphur atoms in compounds **4** and **5**, both perfectly design to produce a chelate ring.

The quenching effect observed could be attributed to an energy transfer quenching of the π^* emissive state through low-lying metal-centered unfilled d-orbitals for Pd^{2+} and Cu^{2+} , and to an intersystem crossing mechanism due to the heavy atom effect.¹⁹

Taking into account the non coordinative counterions used for all metal complexes, (CF_3SO_3^- or BF_4^-) the hypothesis of 2:1 M/L molar ratio could be perfectly predicted as the acetonitrile can acts as a secondary ligand. The stoichiometry of the complexes was obtained through Job's plot. For example, follow the absorption band in the ligand **3** the stoichiometry 1:2 L:M was deduced in the presence of Pd^{2+} (Figure 7).

Figure 7. Job's plot of compound **3** in the presence of Pd²⁺.



The complexation constants of compounds **3-5**, were obtained from the spectroscopic titrations using the Hypspec program.²⁰ The highest constant for Pd²⁺ ions was obtained for compound **3** ($\log K_{\text{ass}} = 11.07 \pm 0.01$) and the value decreased for the other two host systems in the sequential order **3** > **4** > **5**. In the case of Cu²⁺, similar values were obtained for all systems (from $\log K_{\text{ass}} = 10.32 \pm 0.01$ to $\log K_{\text{ass}} = 10.93 \pm 0.01$). For both metal ions, the experimental values suggest the formation of a dinuclear complex. The study of the interaction with fluoride ion indicated a mononuclear species, exhibiting lower constant values in all cases (Table 2).

Table 2. Association constants for benzocrown ethers **3-5** in the presence of F⁻, Pd²⁺ and Cu²⁺ in MeCN/DMSO (99:1).

Cpd.	Guest	log K _{ass} (M:L)
3	F ⁻	3.27± 0.01 (1:1)
	Cu ²⁺	10.32 ± 0.01 (2:1)
	Pd ²⁺	11.07 ± 0.01 (2:1)
4	F ⁻	4.09 ± 0.01 (1:1)
	Cu ²⁺	10.78 ± 0.01 (2:1)
	Pd ²⁺	9.72 ± 0.03 (2:1)
5	F ⁻	4.10 ± 0.02 (1:1)
	Cu ²⁺	10.93 ± 0.01 (2:1)
	Pd ²⁺	10.60 ± 0.01 (2:1)

3. Conclusions

In conclusion, novel imidazo benzocrown ether-based ionophores bearing arylthienyl and bithienyl π -conjugated bridges **3-5** were synthesized in moderate to good yields and evaluated as fluorimetric sensors. Selectivity and sensitivity for Cu²⁺ and Pd²⁺ was observed for all systems. Moreover, systems **4** and **5** proved to be efficient proton and also basic anion sensors, due to changes in emission after deprotonation of the imidazole NH by the fluoride ion.

4. Experimental

4.1. Synthesis general

Progress of the reaction was monitored by thin layer chromatography (0.25 mm thick precoated silica plates: Merck Fertigplatten Kieselgel 60F254), while purification was effected by silica gel column chromatography (Merck Kieselgel 60; 230-400 mesh). NMR spectra were obtained on a Varian Unity Plus Spectrometer at an operating frequency of 300 MHz for ¹H and 75.4 MHz for ¹³C or a Bruker Avance III 400 at an operating frequency of 400 MHz for ¹H and 100.6 MHz for ¹³C, using the solvent peak as internal reference. The solvents are indicated in parenthesis before the chemical shift values (δ relative to TMS). Mps were determined on a Gallenkamp apparatus and are uncorrected. Infrared spectra were

recorded on a BOMEM MB 104 spectrophotometer. Mass spectrometry analyses were performed at the C.A.C.T.I. -Unidad de Espectrometria de Masas of the University of Vigo, Spain. Elemental analysis was carried out on a Leco CHNS-932.

4-Amino-5-nitrobenzo-15-crown-5 was purchased from Aldrich and used as received. The synthesis of 2-(4'-formylphenyl)-thiophene, 2-formyl-5-(4'-formylphenyl)-thiophene and 5,5'-diformyl-2,2'-bithiophene has been described elsewhere.^{6d,13}

4.2. General procedure for the synthesis of compounds 3-5

A solution of 4-amino-5-nitrobenzo-15-crown-5 and the formylated precursor (molar ratio 1:1 for the synthesis of compound **3** and molar ratio 2:1 for the preparation of compounds **4-5**) in DMSO (3 mL) was treated with Na₂S₂O₄ (3 equiv), dissolved in a small volume of water, and heated at 80 °C with stirring for 15 h. The mixture was then cooled to room temperature and the product precipitated during neutralization with NH₄OH 5 M. The precipitate was filtered, washed with water and diethyl ether and dried to give the expected product.

Crown ether **3**. Yellow solid (64%). Mp: 291.5-293.0 °C. IR (Nujol) ν 3443, 2923, 2855, 1639, 1609, 1464, 1377, 1293, 1256, 1219, 1209, 1173, 1139, 1081, 1050, 1008, 983, 945, 849, 826 cm⁻¹. ¹H NMR (DMSO-*d*₆) δ 3.63 (s, 8H, 4xCH₂), 3.82 (s, 4H, 2xCH₂), 4.13 (s, 4H, 2xCH₂), 7.19-7.22 (m, 3H, 4''-H, 4-H and 5-H), 7.67 (dd, 1H, *J*=4.5 and 0.9 Hz, 5''-H), 7.71 (dd, 1H, *J*=2.7 and 0.9 Hz, 3''-H), 7.93 (d, 2H, *J*=8.7 Hz, 2'-H and 6'-H), 8.13 (d, 2H, *J*=8.7 Hz, 3'-H and 5'-H). ¹³C NMR (DMSO-*d*₆) δ 68.71 (2xCH₂), 68.99 (2xCH₂), 69.71 (2xCH₂), 70.52 (2xCH₂), 97.93 (C4 and C5), 124.12 (C4'), 125.42 (C3''), 126.04 (C2' and C6'), 127.39 (C5''), 127.61 (C3' and C5'), 128.14 (C3a and C5a), 128.97 (C4''), 136.53 (C1'), 141.99 (C2''), 147.06 (C2), 148.16 (C4a and C4b). MS (FAB) *m/z* (%): 467 ([M+H]⁺, 60), 466 (M⁺, 17), 308 (13), 307 (40), 289 (17), 155 (32), 154 (100). HRMS: (FAB) *m/z* (%) for C₂₅H₂₇N₂O₅S; calcd 467.1641; found 467.1654.

Crown ether **4**. Brown solid (78%). Mp: 246.5-248.9 °C. IR (Nujol) ν 3449, 2930, 2842, 1633, 1593, 1467, 1378, 1284, 1121, 1053, 938, 844, 811, 722 cm⁻¹. ¹H NMR (DMSO-*d*₆) 3.65 (s, 16H, 4xCH₂), 3.82 (s, 8H, 2xCH₂), 4.11 (s, 8H, 2xCH₂), 7.14 (s, 2H, 4-H and 5-H), 7.18 (s, 2H, 4-H and 5-H), 7.71 (d, 1H, *J*=4.0 Hz, 3''-H), 7.87 (d, 2H, *J*=8.8 Hz, 2'-H and 6'-H), 7.92 (d, 1H, *J*=4.0 Hz, 4''-H), 8.27 (d, 2H, *J*=8.4 Hz, 3'-H and 5'-H), 13.45 (br s, 1H, NH). ¹³C NMR (DMSO-*d*₆) δ 68.46 (4xCH₂), 68.52 (4xCH₂), 69.13 (4xCH₂), 69.65 (2xCH₂), 69.73 (2xCH₂), 125.40, 125.69, 126.89, 127.22, 128.62, 133.76, 134.20, 143.41, 145.38, 145.84,

146.22, 148.81. MS (FAB) m/z (%): 795 ($[M+Na]^+$, 100), 773 ($[M+H]^+$, 21), 636 (12), 517 (10). HRMS: (FAB) m/z (%) for $C_{40}H_{45}N_4O_{10}S$; calcd 773.2856; found 773.2866.

Crown ether **5**. Brown solid (53%). Mp: 260.5-263.0 °C. IR (KBr) ν 3451, 3078, 2922, 2855, 1655, 1490, 1439, 1345, 1297, 1226, 1130, 1053, 943, 799 cm^{-1} . 1H NMR (DMSO- d_6) 3.63 (s, 16H, 8xCH₂), 3.82 (s, 8H, 4xCH₂), 4.08 (s, 8H, 4xCH₂), 6.99 (s, 2H, 2x4-H), 7.17 (s, 2H, 2x5-H), 7.45 (d, 2H, $J=4.0$ Hz, 2x3'-H), 7.68 (d, 2H, $J=4.0$ Hz, 2x4'-H), 12.75 (s, 2H, 2xNH). ^{13}C NMR (DMSO- d_6) δ 68.88 (8xCH₂), 69.79 (4xCH₂), 70.37 (4xCH₂), 95.88 (2xC4 and 2xC5), 125.35 (2xC3'), 126.30 (2xC4'), 127.81 (2xC3a and 2xC5a), 133.27 (2xC2'), 136.86 (2xC5'), 144.92 (2xC2), 146.04 (2xC4a and 2xC4b). MS (FAB) m/z (%): 779 ($[M+H]^+$, 10), 501 (34), 307 (34), 282 (22), 155 (32), 154 (100). HRMS: (FAB) m/z (%) for $C_{38}H_{43}N_4O_{10}S_2$; calcd 779.2421; found 779.2394.

4.3. Spectrophotometric and spectrofluorimetric titrations of compounds 3-5

UV-vis absorption spectra (220 – 800 nm) were obtained using a Perkin Elmer lambda 35 spectrophotometer and fluorescence emission on a Perkin Elmer LS45. The linearity of the fluorescence emission vs. concentration was checked in the concentration range used (10^{-4} - 10^{-6} M). A correction for the absorbed light was performed when necessary. All spectrofluorimetric titrations were performed as follows: stock solutions of compounds **3-5** (ca. 10^{-3} M) were prepared in a 50 mL volumetric flask and diluting to the mark with MeCN/DMSO (99:1) UVA-sol. The titration solutions were prepared by appropriate dilution of stock solutions. Titrations of compounds **3-5** were carried out by addition of microliter amounts of standard solutions of the ions (cations or anions) in acetonitrile.

The organic solvents were purified by standard methods.²¹ CH₄SO₃, Bu₄NOH, NaBF₄, KBF₄, LiBF₄, Ca(CF₃SO₃)₂, Zn(BF₄)₂, Cu(BF₄)₂, Ni(BF₄)₂, Pd(BF₄)₂, Hg(CF₃SO₃)₂, and Bu₄NF were purchased from Strem Chemicals, Sigma Aldrich and Solchemar. All were used without previous purification. CH₄SO₃ was used to change the acidity conditions of the MeCN/DMSO solutions. Luminescence quantum yields were measured using a solution of quinine sulfate in sulfuric acid (0.1M) as standard [$\phi_F = 0.54$].²² All measurements were conducted at 298 K.

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Captions

Table 1. Yields, IR, ¹H NMR, UV-vis absorption and fluorescence data for imidazo-benzocrown ethers **3-5**.

^a For the NH stretching band (recorded in nujol).

^b For the NH stretching band (recorded in KBr).

^c For the NH proton of the imidazole ring (in DMSO-d₆).

^d in MeCN/DMSO (99:1) solution.

Table 2. Association constants for benzocrown ethers **3-5** in the presence of F⁻, Pd²⁺ and Cu²⁺ in MeCN/DMSO (99:1).

Figure 1. Absorption and emission spectra of compounds **3-5** in MeCN/DMSO (99:1), T= 298 K, [**3**] = 5.0×10⁻⁷ M, [**4**] = 5.5×10⁻⁷ M, [**5**] = 4.3 ×10⁻⁷ M. (λ_{exc3} = 360 nm, λ_{exc4} = 395 nm; λ_{exc5} = 415 nm).

Figure 2. Spectrofluorimetric titration of compounds **4** (A) and **5** (B) in MeCN/DMSO (99:1) solutions as a function of added CH₄O₃S. The insets show the normalized fluorescence intensity at 500 nm and 535 nm, respectively ([**4**] = 4.30×10⁻⁷ M, λ_{exc} = 395 nm; [**5**] = 5.10×10⁻⁷ M, λ_{exc} = 415 nm).

Figure 3. Absorption and fluorescence spectra of MeCN/DMSO (99:1) solutions of **3** (A), **4** (B) and **5** (C) and fluorescence titration as a function of Bu₄NF added. The insets show the normalized fluorescence intensity at 447 nm (**3**), 496 nm (**4**) and 568 nm (**5**) ([**3**] = 5.13×10⁻⁷ M, λ_{exc} = 360 nm; [**4**] = 4.30×10⁻⁷ M, λ_{exc} = 395 nm). [**5**] = 5.10×10⁻⁷ M, λ_{exc} = 415 nm).

Figure 4. Absorption and fluorescence spectra of MeCN/DMSO (99:1) solutions of **3** (A), **4** (B) and **5** (C) after addition of 2000 equivalents of NaBF₄; LiBF₄; KBF₄; Ca(CF₃SO₃)₂ and Zn(CF₃SO₃)₂ in CH₃CN. ([**3**] = 5.13×10⁻⁷ M, λ_{exc} = 360 nm; [**4**] = 4.30×10⁻⁷ M, λ_{exc} = 395 nm; [**5**] = 5.10×10⁻⁷ M, λ_{exc} = 415 nm).

Figure 5. Absorption spectra of acetonitrile-DMSO (99:1) solutions of **3** (A), **4** (B) and **5** (C) and fluorescence titration as a function of added Pd(CH₃CN)₄BF₄. The insets show the normalized fluorescence intensity at 447 nm (**3**), 500 nm (**4**) and 568 nm (**5**). (T= 298 K; [**3**] = 5.13.10⁻⁷ M, λ_{exc} = 360 nm; [**4**] = 5.10.10⁻⁷ M, λ_{exc} = 395 nm; [**5**] = 4.30.10⁻⁷ M, λ_{exc} = 415 nm).

Figure 6. Absorption and fluorescence spectra of MeCN/DMSO (99:1) solutions of **3** (A), **4** (B) and **5** (C) as a function of added Cu(CF₃SO₃)₂. The insets show the normalized fluorescence intensity at 447 nm (**3**), 500 nm (**4**) and 568 nm (**5**). ([**3**] = 5.13×10⁻⁷ M, λ_{exc} = 360 nm; [**4**] = 4.30×10⁻⁷ M, λ_{exc} = 395 nm, [**5**] = 5.10×10⁻⁷ M, λ_{exc} = 415 nm).

Figure 7. Job's plot of compound **3** in the presence of Pd²⁺.

Scheme 1. Synthesis of imidazo-benzocrown ether derivatives **3-5**.

Research Highlights

- 1- Synthesis of novel imidazo-benzo-15-crown-5 ethers bearing arylthienyl and bithienyl π -bridges.
- 2- Characterization of the absorption and emission properties of imidazo-benzo-15-crown-5 ethers.
- 3- Thienyl derivatives of benzo-15-crown-5 ethers as selective fluorescent chemosensors for Pd^{2+} and Cu^{2+} .
- 4- *bis*-Substituted imidazo-crown ethers as selective fluorimetric sensors for the fluoride ion.